

APPENDIX II
CLEAN VERSION OF THE ENTIRE SET OF
PENDING CLAIMS AS AMENDED IN THIS COMMUNICATION
PURSUANT TO 37 CFR § 1.121 (c)(1)(i)

The following is a list of the all claims as they appear following entry of the amendments set out in the Response to the pending non-final Office Action.

1. (Amended) A method for producing recombinant mini-Adenovirus comprising:
 - a) providing:
 - i) a first recombinant vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest;
 - 3) adenovirus packaging sequence linked to one of said inverted terminal repeats; and
 - 4) an adeno-associated virus terminal repeat sequence operably linked to said 3' end of said nucleotide sequence of interest,
 - wherein said first vector lacks a second adeno-associated virus terminal repeat sequence, and lacks one or more adenovirus early gene region selected from E1, E2, E3, and E4 gene region; and
 - ii) a cell capable of expressing said one or more adenovirus early gene which is lacking from said first vector;
- b) introducing said first vector into said cell to produce a transformed cell; and
- c) culturing said transformed cell under conditions such that a second vector is produced, said second vector selected from:

- i) a vector, comprising in operable combination:
 - 1) adeno-associated virus terminal repeat-DD sequence;
 - 2) first and second inverted copies of a nucleotide sequence of interest flanking said adeno-associated virus terminal repeat-DD sequence;
 - 3) left and right inverted terminal repeats of adenovirus flanking said first and second inverted copies of said nucleotide sequence of interest; and
 - 4) an adenovirus packaging sequence linked to one of said inverted terminal repeats, and
- ii) a vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest; and
 - 3) an adenovirus packaging sequence linked to one of said inverted terminal repeats.

2. (Amended) The method for producing recombinant mini-Adenovirus of Claim 1, wherein said cell is capable of expressing one or more Rep proteins, and said culturing results in expression of said one or more Rep proteins.

3. (Amended) The method for producing recombinant mini-Adenovirus of Claim 1, wherein said second vector is encapsidated.

4. (Amended) The method for producing recombinant mini-Adenovirus of Claim 3, further comprising d) recovering said encapsidated second vector.

5. (Amended) The method for producing recombinant mini-Adenovirus of Claim 4, further comprising e) purifying said recovered encapsidated second vector.

11. (Amended) The method for producing recombinant mini-Adenovirus of Claim 2, wherein expression of one or more Rep proteins is inducible.

12. (Amended) A method for producing recombinant mini-Adenovirus comprising:

- a) providing:
 - i) a first recombinant vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest;
 - 3) adenovirus packaging sequence linked to one of said inverted terminal repeats; and
 - 4) an adeno-associated virus terminal repeat sequence operably linked to said 3' end of said nucleotide sequence of interest,wherein said first vector lacks a second adeno-associated virus terminal repeat sequence, and lacks one or more adenovirus early gene region selected from E1, E2, and E4 gene region;
 - ii) a cell capable of expressing one or more Rep proteins; and
 - iii) helper adenovirus;
- b) introducing said first vector and genome of said helper adenovirus into said cell to produce a transformed cell; and
- c) culturing said transformed cell under conditions such that said transformed cell expresses said one or more Rep proteins, and a second vector is produced, said second vector selected from:

- i) a vector, comprising in operable combination:
 - 1) adeno-associated virus terminal repeat-DD sequence;
 - 2) first and second inverted copies of a nucleotide sequence of interest flanking said adeno-associated virus terminal repeat-DD sequence;
 - 3) left and right inverted terminal repeats of adenovirus flanking said first and second inverted copies of said nucleotide sequence of interest; and
 - 4) an adenovirus packaging sequence linked to one of said inverted terminal repeats, and
- ii) a vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest; and
 - 3) an adenovirus packaging sequence linked to one of said inverted terminal repeats.

13. (Amended) The method for producing recombinant mini-Adenovirus of Claim 12, wherein said cell lacks expression of said one or more adenovirus early gene region which is lacking from said first vector.

14. (Amended) A method for producing recombinant mini-Adenovirus comprising:

- a) providing:
 - i) a first recombinant vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;

- 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest;
- 3) adeno-associated virus terminal repeat sequence operably linked to said 3' end of said nucleotide sequence of interest,

wherein said first vector lacks a second adeno-associated virus terminal repeat sequence, and lacks one or more adenovirus early gene region selected from E1, E2, and E4 gene region;

- ii) a cell capable of expressing said one or more adenovirus early gene region selected from E1, E2, and E4 gene region;
 - iii) a cell capable of expressing said one or more adenovirus early gene which is lacking from said first vector; and
 - iii) adeno-associated virus;
- b) introducing said first vector and genome of said adeno-associated virus into said cell to produce a transformed cell; and
- c) culturing said transformed cell under conditions such that a second vector is produced, said second vector selected from:
- i) a vector, comprising in operable combination:
 - 1) adeno-associated virus terminal repeat-DD sequence;
 - 2) first and second inverted copies of a nucleotide sequence of interest flanking said adeno-associated virus terminal repeat-DD sequence;
 - 3) left and right inverted terminal repeats of adenovirus flanking said first and second inverted copies of said nucleotide sequence of interest; and
 - 4) an adenovirus packaging sequence linked to one of said inverted terminal repeats, and
 - ii) a vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;

- 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest; and
- 3) an adenovirus packaging sequence linked to one of said inverted terminal repeats.

15. (Amended) A method for producing recombinant mini-Adenovirus comprising:

- a) providing:
 - i) a first recombinant vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest;
 - 3) adenovirus packaging sequence linked to one of said inverted terminal repeats; and
 - 4) an adeno-associated virus terminal repeat sequence operably linked to said 3' end of said nucleotide sequence of interest,wherein said first vector lacks a second adeno-associated virus terminal repeat sequence, and lacks adenovirus E3 early gene region; and
 - ii) a cell;
- b) introducing said first vector into said cell to produce a transformed cell; and
- c) culturing said transformed cell under conditions such that a second vector is produced, said second vector selected from:
 - i) a vector, comprising in operable combination:
 - 1) adeno-associated virus terminal repeat-DD sequence;

- 2) first and second inverted copies of a nucleotide sequence of interest flanking said adeno-associated virus terminal repeat-DD sequence;
 - 3) left and right inverted terminal repeats of adenovirus flanking said first and second inverted copies of said nucleotide sequence of interest; and
 - 4) an adenovirus packaging sequence linked to one of said inverted terminal repeats, and
- ii) a vector, comprising in operable combination:
- 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest; and
 - 3) an adenovirus packaging sequence linked to one of said inverted terminal repeats.

16. (Amended) The method for producing recombinant mini-Adenovirus of Claim 15, wherein said cell is capable of expressing one or more of Rep proteins, and said culturing results in expression of said one or more Rep proteins.

17. (Amended) A method for producing recombinant mini-Adenovirus comprising:

- a) providing:
 - i) a first recombinant vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest;
 - 3) adenovirus packaging sequence linked to one of

said inverted terminal repeats; and

- 4) an adeno-associated virus terminal repeat sequence operably linked to said 3' end of said nucleotide sequence of interest,

wherein said first vector lacks a second adeno-associated virus terminal repeat sequence, and wherein said nucleotide sequence of interest in said first vector comprises adeno-associated virus Rep gene region; and

- ii) a cell;
- b) introducing said first vector into said cell to produce a transformed cell; and
- c) culturing said transformed cell under conditions such that said transformed cell expresses one or more Rep proteins, and a second vector is produced, said second vector selected from:
 - i) a vector, comprising in operable combination:
 - 1) adeno-associated virus terminal repeat-DD sequence;
 - 2) first and second inverted pieces of a nucleotide sequence of interest flanking said adeno-associated virus terminal repeat-DD sequence;
 - 3) left and right inverted terminal repeats of adenovirus flanking said first and second inverted copies of said nucleotide sequence of interest; and
 - 4) an adenovirus packaging sequence linked to one of said inverted terminal repeats, and
 - ii) a vector, comprising in operable combination:
 - 1) a nucleotide sequence of interest having a 5' end and a 3' end;
 - 2) left and right inverted terminal repeats of adenovirus flanking said nucleotide sequence of interest; and

- 3) an adenovirus packaging sequence linked to one of said inverted terminal repeats.

18. (Amended) The method for producing recombinant mini-Adenovirus of Claim 17, wherein said first vector lacks one or more adenovirus early gene region selected from E1, E2, and E4 gene region, and said cell is capable of expressing said adenovirus early gene region which is lacking from said first vector.

19. (Amended) The method for producing recombinant mini-Adenovirus of Claim 17, wherein said first vector lacks adenovirus E3 gene region.